

SEQUENCE LISTING

<110 Houghton, Michael Choo, Oui-Lim Kuo, George

<120> Hepatitis C virus protease

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<141> 2001-06-18

<150> 09/253,675

<151> 1999-02-18

<150> 08/709,177

<151> 1996-09-06

<150> 08/440,548

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<150> 08/350,884

<151> 1994-12-06

<150> 07/680,296

<151> 1991-04-04

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Phe Leu Ala Thr Cys Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly
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Ala Gly Thr Arg Thr Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met
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Tyr Thr Asn Val Asp Gln Asp Leu Val Gly Trp Pro Ala Ser Gln Gly
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Thr Arg Ser Leu Thr Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu
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Arg Gly Ser Leu Leu Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser
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Ser Gly Gly Pro Leu Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe
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<223> Elastase protease
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Gly Tyr Asp Ile Ala Leu Leu
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Gly Asp Ser Gly Gly Pro Leu
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Arg Gly Leu Leu Gly Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys
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Asn Gln Val Glu Gly Glu Val Gln Ile Val Ser Thr Ala Ala Gln Thr
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Phe Leu Ala Thr Cys Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly
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Ala Gly Thr Arg Thr Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met
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Tyr Thr Asn Val Asp Gln Asp Leu Val Gly Trp Pro Ala Ser Gln Gly
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Thr Arg Ser Leu Thr Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu
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                            120
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Val Thr Arg His Ala Asp Val Ile Pro Val Arg Arg Gly Asp Ser
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Arg Gly Ser Leu Leu Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser
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                                         155
                    150
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Ser Gly Gly Pro Leu Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe
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                                     170
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Arg Ala Ala Val Cys Thr Arg Gly Val Ala Lys Ala Val Asp Phe Ile
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Gln Met Glu Thr Lys Leu Ile Thr Trp Gly Ala Asp Thr Ala Ala Cys
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Gly Asp Ile Ile Asn Gly Leu Pro Val Ser Ala Arg Arg Gly Arg Glu
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Ile Leu Leu Gly Pro Ala Asp Gly Met Val Ser Lys Gly Trp Arg Leu
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Leu Ala Pro Ile Thr Ala Tyr Ala Gln Gln Thr Arg Gly Leu Leu Gly
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Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn Gln Val Glu Gly
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Glu Val Gln Ile Val Ser Thr Ala Ala Gln Thr Phe Leu Ala Thr Cys
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Ile Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Thr Arg
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Thr Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met Tyr Thr Asn Val
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Asp Gln Asp Leu Val Gly Trp Pro Ala Ser Gln Gly Thr Arg Ser Leu
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 Thr Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu Val Thr Arg His
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Ala Asp Val Ile Pro Val Arg Arg Gly Asp Ser Arg Gly Ser Leu
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Leu Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro
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Leu Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe Arg Ala Ala Val
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Cys Thr Arg Gly Val Ala Lys Ala Val Asp Phe Ile Pro Val Glu Asn
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Leu Glu Thr Thr Met Arg Ser Pro Val Phe Thr Asp Asn Ser Ser Pro
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Pro Val Val Pro Gln Ser Phe Gln Val Ala His Leu His Ala Pro Thr
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Gly Ser Gly Lys Ser Thr Lys Val Pro Ala Ala
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Gly Asp Ile Ile Asn Gly Leu Pro Val Ser Ala Arg Arg Gly Arg Glu
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Ile Leu Leu Gly Pro Ala Asp Gly Met Val Ser Lys Gly Trp Arg Leu
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Leu Ala Pro Ile Thr Ala Tyr Ala Gln Gln Thr Arg Gly Leu Leu Gly
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Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn Gln Val Glu Gly
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                                 105
                                                     110
Glu Val Gln Ile Val Ser Thr Ala Ala Gln Thr Phe Leu Ala Thr Cys
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                            120
                                                 125
Ile Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Thr Arg
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Thr Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met Tyr Thr Asn Val
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Asp Gln Asp Leu Val Gly Trp Pro Ala Ser Gln Gly Thr Arg Ser Leu
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Gln Met Glu Thr Lys Leu Ile Thr Trp Gly Ala Asp Thr Ala Ala Cys
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Gly Asp Ile Ile Asn Gly Leu Pro Val Ser Ala Arg Arg Gly Arg Glu
                         55
Ile Leu Leu Gly Pro Ala Asp Gly Met Val Ser Lys Gly Trp Arg Leu
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                     70
                                         75
                                                             80
Leu Ala Pro Ile Thr Ala Tyr Ala Gln Gln Thr Arg Gly Leu Leu Gly
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                                     90
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Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn Gln Val Glu Gly
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                                 105
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Glu Val Gln Ile Val Ser Thr Ala Ala Gln Thr Phe Leu Ala Thr Cys
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                             120
                                                 125
Ile Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Thr Arg
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Thr Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met Tyr Thr Asn Val
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Asp Gln Asp Leu Val Gly Trp Pro Ala Ser Gln Gly Thr Arg Ser Leu
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                                     170
                                                         175
Thr Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu Val Thr Arg His
            180
                                 185
                                                     190
Ala Asp Val Ile Pro Val Arg Arg Gly Asp Ser Arg Gly Ser Leu
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                            200
                                                 205
Leu Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro
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                        215
                                             220
Leu Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe Arg Ala Ala Val
225
                    230
                                         235
Cys Thr Arg Gly Val Ala Lys Ala Val Asp Phe Ile Pro Val Glu Asn
                245 250
Leu Glu Thr Thr Met Arg Ser Pro Val Phe Thr Asp Asn Ser Ser Pro
            260
                                265
                                                     270
Pro Val Val Pro Gln Ser Phe Gln Val Ala His Leu His Ala Pro Thr
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                            280
                                                 285
Gly Ser Gly Lys Ser Thr Lys Val Pro Ala Ala
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                        295
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                                                                       120
atcacgtggg gggcagatac cgccgcgtgc ggtgacatca tcaacggctt gcctgtttcc
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gcccgcaggg gccgggagat actgctcggg ccagccgatg gaatggtctc caagggttgg
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                                                                      480
cacggggccg gaacgaggac catcgcgtca cccaagggtc ctgtcatcca gatgtatacc
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                                                                     1680
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gcaccggggg agcgccctcc cggcatgttc gactcgtccg tcctctgtga gtgctatgac
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gcaggctgtg cttggtatga gctcacgccc gccgagacta cagttaggct acgagcgtac
                                                                     1860
atgaacaccc cggggcttcc cgtgtgccag gaccatcttg aattttggga gggcgtcttt
                                                                     1920
acaggeetea eteatataga tgeecaettt etateeeaga caaageagag tggggagaae
                                                                     1980
cttccttacc tggtagcgta ccaagccacc gtgtgcgcta gggctcaagc ccctcccca
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Ile Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Thr Arg Thr Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met Tyr Thr Asn Val Asp Gln Asp Leu Val Gly Trp Pro Ala Ser Gln Gly Thr Arg Ser Leu Thr Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu Val Thr Arg His 190· Ala Asp Val Ile Pro Val Arg Arg Gly Asp Ser Arg Gly Ser Leu Leu Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro Leu Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe Arg Ala Ala Val Cys Thr Arg Gly Val Ala Lys Ala Val Asp Phe Ile Pro Val Glu Asn Leu Glu Thr Thr Met Arg Ser Pro Val Phe Thr Asp Asn Ser Ser Pro Pro Val Val Pro Gln Ser Phe Gln Val Ala His Leu His Ala Pro Thr Gly Ser Gly Lys Ser Thr Lys Val Pro Ala Ala Tyr Ala Ala Gln Gly Tyr Lys Val Leu Val Leu Asn Pro Ser Val Ala Ala Thr Leu Gly Phe Gly Ala Tyr Met Ser Lys Ala His Gly Ile Asp Pro Asn Ile Arg Thr Gly Val Arg Thr Ile Thr Thr Gly Ser Pro Ile Thr Tyr Ser Thr Tyr Gly Lys Phe Leu Ala Asp Gly Gly Cys Ser Gly Gly Ala Tyr Asp Ile Ile Ile Cys Asp Glu Cys His Ser Thr Asp Ala Thr Ser Ile Leu Gly Ile Gly Thr Val Leu Asp Gln Ala Glu Thr Ala Gly Ala Arg Leu Val Val Leu Ala Thr Ala Thr Pro Pro Gly Ser Val Thr Val Pro His Pro Asn Ile Glu Glu Val Ala Leu Ser Thr Thr Gly Glu Ile Pro Phe Tyr Gly Lys Ala Ile Pro Leu Glu Val Ile Lys Gly Gly Arg His Leu Ile Phe Cys His Ser Lys Lys Lys Cys Asp Glu Leu Ala Ala Lys Leu Val Ala Leu Gly Ile Asn Ala Val Ala Tyr Tyr Arg Gly Leu Asp Val Ser Val Ile Pro Thr Ser Gly Asp Val Val Val Ala Thr Asp Ala Leu Met Thr Gly Tyr Thr Gly Asp Phe Asp Ser Val Ile Asp Cys Asn Thr Cys Val Thr Gln Thr Val Asp Phe Ser Leu Asp Pro Thr Phe Thr Ile Glu Thr Ile Thr Leu Pro Gln Asp Ala Val Ser Arg Thr Gln Arg Arg

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530
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Gly Arg Thr Gly Arg Gly Lys Pro Gly Ile Tyr Arg Phe Val Ala Pro
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Gly Glu Arg Pro Pro Gly Met Phe Asp Ser Ser Val Leu Cys Glu Cys
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Tyr Asp Ala Gly Cys Ala Trp Tyr Glu Leu Thr Pro Ala Glu Thr Thr
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Val Arg Leu Arg Ala Tyr Met Asn Thr Pro Gly Leu Pro Val Cys Gln
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Asp His Leu Glu Phe Trp Glu Gly Val Phe Thr Gly Leu Thr His Ile
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Asp Ala His Phe Leu Ser Gln Thr Lys Gln Ser Gly Glu Asn Leu Pro
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Tyr Leu Val Ala Tyr Gln Ala Thr Val Cys Ala Arg Ala Gln Ala Pro
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                                     650
Pro Pro Ser Trp Asp Gln Met Trp Lys Cys Leu Ile Arg Leu Lys Pro
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Thr Leu His Gly Pro Thr Pro Leu Leu Tyr Arg Leu Gly Ala
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ctttacctgg tcacgaggca cgccgatgtc attcccgtgc gccggcgggg tgatagcagg
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ccgaattc
<210> 72
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            20
                                 25
                                                      30
Tyr His Gly Ala Gly Thr Arg Thr Ile Ala Ser Pro Lys Gly Pro Val
        35
Ile Gln Met Tyr Thr Asn Val Asp Gln Asp Leu Val Gly Trp Pro Ala
    50
                         55
Ser Gln Gly Thr Arg Ser Leu Thr Pro Cys Thr Cys Gly Ser Ser Asp
65
                     70
                                         75
                                                              80
Leu Tyr Leu Val Thr Arg His Ala Asp Val Ile Pro Val Arg Arg Arg
                85
                                     90
                                                          95
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Lys Gly Ser Ser Gly Gly Pro Leu Pro Asn
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atgaggtccc cggtgttcac ggataactcc tctccaccag tagtgcccca gagcttccag
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gtggctcacc tccatgctcc ccgaattc
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<210> 74
<211> 69
<212> PRT
<213> Hepatitis C virus
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            20
                                 25
                                                     30
Pro Val Glu Asn Leu Glu Thr Thr Met Arg Ser Pro Val Phe Thr Asp
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His Ala Pro Arg Ile
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<213> Hepatitis C virus
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gccgcggtgt gcacccgtgg agtggctaag gcggtggact ttatccctgt ggagaaccta
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<213> Hepatitis C virus
<400> 76
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1
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Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro Leu
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Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe Arg Ala Ala Val Cys
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Thr Arg Gly Val Ala Lys Ala Val Asp Phe Ile Pro Val Glu Asn Leu
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Phe Ser Gln Met Glu Thr Lys Leu Ile Thr Trp Gly Ala Asp Thr Ala
Ala Cys Gly Asp Ile Ile Asn Gly Leu Pro Val Ser Ala Arg Arg Gly
                        55
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Arg Glu Ile Leu Leu Gly Pro Ala Asp Gly Met Val Ser Lys Gly Trp
65
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Arg Leu Leu Ala Pro Ile Thr Ala Tyr Ala Gln Gln Thr Arg Gly Leu
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                                    90
                                                        95
Leu Gly Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn Gln Val
            100
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Glu Gly Glu Val Gln Ile Val Ser Thr Ala Ala Gln Thr Phe Leu Ala
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Thr Cys Ile Asn Gly Val Cys Trp Pro Asn
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<212> DNA
<213> Hepatitis C virus
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gctgtctccc gcactcaacg tcggggcagg actggcaggg ggaagccagg catctacaga
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ccgaattc
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Asp Cys Asn Thr Cys Val Thr Gln Thr Val Asp Phe Ser Leu Asp Pro
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                             40
                                                 45
Thr Phe Thr Ile Glu Thr Ile Thr Leu Pro Gln Asp Ala Val Ser Arg
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                                             60
Thr Gln Arg Arg Gly Arg Thr Gly Arg Gly Lys Pro Gly Ile Tyr Arg
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                                         75
Phe Val Ala Pro Gly Glu Arg Pro Ser Gly Met Phe Asp Ser Ser Val
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                                     90
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Leu Cys Glu Cys Pro Asn
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<212> PRT

<213> Hepatitis C virus

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720

780

816

<210> 84 <211> 272

<212> PRT

<213> Hepatitis C virus

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Ser Thr Lys Val Pro Ala Ala Tyr Ala Ala Gln Gly Tyr Lys Val Leu
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Val Leu Asn Pro Ser Val Ala Ala Thr Leu Gly Phe Gly Ala Tyr Met
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Ser Lys Ala His Gly Ile Asp Pro Asn Ile Arg Thr Gly Val Arg Thr
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Ile Thr Thr Gly Ser Pro Ile Thr Tyr Ser Thr Tyr Gly Lys Phe Leu
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Ala Asp Gly Gly Cys Ser Gly Gly Ala Tyr Asp Ile Ile Cys Asp
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Glu Cys His Ser Thr Asp Ala Thr Ser Ile Leu Gly Ile Gly Thr Val
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                         135
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Leu Asp Gln Ala Glu Thr Ala Gly Ala Arg Leu Val Val Leu Ala Thr
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Ala Thr Pro Pro Gly Ser Val Thr Val Pro His Pro Asn Ile Glu Glu
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Val Ala Leu Ser Thr Thr Gly Glu Ile Pro Phe Tyr Gly Lys Ala Ile
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                                                     190
Pro Leu Glu Val Ile Lys Gly Gly Arg His Leu Ile Phe Cys His Ser
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                             200
                                                 205
Lys Lys Cys Asp Glu Leu Ala Ala Lys Leu Val Ala Leu Gly Ile
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                                             220
Asn Ala Val Ala Tyr Tyr Arg Gly Leu Asp Val Ser Val Ile Pro Thr
225
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Ser Gly Asp Val Val Val Ala Thr Asp Ala Leu Met Thr Gly Tyr
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                                25
                                                    30
Trp Gly Ser Ile Lys Gly Leu Thr Glu Gly Leu His Gly Phe His Val
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480

540

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660

720

780

840

900

960

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40

His Glu Phe Gly Asp Asn Thr Ala Gly Cys Thr Ser Pro Gly Pro His Phe Asn Pro Leu Ser Arg Lys His Gly Gly Pro Lys Asp Glu Glu Arg His Val Gly Asp Leu Gly Asn Val Thr Ala Asp Lys Asp Gly Val Ala Asp Val Ser Ile Glu Asp Ser Val Ile Ser Leu Ser Gly Asp His Cys Ile Ile Gly Arg Thr Leu Val Val His Glu Lys Ala Asp Asp Leu Gly Lys Gly Gly Asn Glu Glu Ser Thr Lys Thr Gly Asn Ala Gly Ser Arg Leu Ala Cys Gly Val Ile Gly Ile Arg Ile Arg Gly Thr Tyr Val Tyr Asn His Leu Thr Pro Leu Arg Asp Trp Ala His Asn Gly Leu Arg Asp Leu Ala Val Ala Val Glu Pro Val Val Phe Ser Gln Met Glu Thr Lys Leu Ile Thr Trp Gly Ala Asp Thr Ala Ala Cys Gly Asp Ile Ile Asn Gly Leu Pro Val Ser Ala Arg Arg Gly Arg Glu Ile Leu Leu Gly Pro Ala Asp Gly Met Val Ser Lys Gly Trp Arg Leu Leu Ala Pro Ile Thr Ala Tyr Ala Gln Gln Thr Arg Gly Leu Leu Gly Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn Gln Val Glu Gly Glu Val Gln Ile Val 265... Ser Thr Ala Ala Gln Thr Phe Leu Ala Thr Cys Ile Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Thr Arg Thr Ile Ala Ser Pro Lys Gly Pro Val İle Gln Met Tyr Thr Asn Val Asp Gln Asp Leu Val Gly Trp Pro Ala Ser Gln Gly Thr Arg Ser Leu Thr Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu Val Thr Arg His Ala Asp Val Ile Pro Val Arg Arg Gly Asp Ser Arg Gly Ser Leu Leu Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro Leu Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe Arg Ala Ala Val Cys Thr Arg Gly Val Ala Lys Ala Val Asp Phe Ile Pro Val Glu Asn Leu Glu Thr Thr Met Arg Ser Pro Val Phe Thr Asp Asn Ser Ser Pro Pro Val Val Pro Gln Ser Phe Gln Val Ala His Leu His Ala Pro Thr Gly Ser Gly Lys Ser Thr Lys Val Pro Ala Ala Tyr Ala Ala Gln Gly Tyr Lys Val Leu Val

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465
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Lys Ala His Gly Ile Asp Pro Asn Ile Arg Thr Gly Val Arg Thr Ile
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Cys His Ser Thr Asp Ala Thr Ser Ile Leu Gly Ile Gly Thr Val Leu
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Thr Pro Pro Gly Ser Val Thr Val Pro His Pro Asn Ile Glu Glu Val
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Pro Gln Asp Ala Val Ser Arg Thr Gln Arg Arg Gly Arg Thr Gly Arg
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Gly Met Phe Asp Ser Ser Val Leu Cys Glu Cys Tyr Asp Ala Gly Cys
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